

Fig. 1A

Sequence ID No. 4

CGGGAGAATAGTGCACCAAGGGATGCCCCGTGAAATATTAATTAAACGTTTTTTAAGAACA -101
TCATCAAACCCGGGCCCCATCATGAAGGAATAACAAGGCCTTCGAAAAGTATGGGAAACT -41
GGTCGGCAGGACATCAGCATTATTAATTCTAGGAAACTCATTATGGATAACAAGGAAACT 18
M D N K E T 6
AACGGAGAGCTAGAGCAGTCTGATGAGGCCGATCCGTCCGGTCAAAACCTTGATGATGGG 78
N G E L E Q S D E A D P S G Q N L D D G 26
GAAACCGATAGCAAACAAGAAGAGAATCTCATCAACGTTAGCCCGCCAAAAACACCGCCA 138
E T D S K Q E E N L I N V S P P K T P P 45
GGTCCTCCTCCTCCTCTAAAGAATGGAGGAAGGGGTCAGAAACCGCCCAAAATCCCAATA 198
G P P P P L K N G G R G Q K P P K I P I 66
TGTCATCAAAATGGAAAGCTCCCCAAGGAAGTTGAATGGACAGAAGACAGAGGGCGAAGAC 258
C H Q N G K L P K E V E W T E D R G E D 86
AGAAAGGATAGTCTCACTCTTCAATCAAAGCTAGATCACGGGGCATAACACGGATGAGAAA 318
R K D S L T L Q S K L D H G A Y T D E K 106
CAGGATCTTCTAACATATCTTGACCGTCACGGCATCAACAGTCCAGTCAAGCTAACACCA 378
Q D L L T Y L D R H G I N S P V K L T P 126
GATGAAACTGGAGGGAGCAGTGCTTTGGATATTCTTGGGATTATTGAAGAGAGGGGACACT 438
D E T G G S S A L D I L G I I E E R D T 146
GGTGCCTAGGCTCTGATCCCTCATCCACTATGCAGGCCATGGCTAAACCTGTAGGCTTT 498
G A L G S D P S S T M Q A M A K P V G F 166
CTGCAGAGGCAGCTATGGACTGTCCTCCAACCTTCAGACAATAGACTCTCCATGAAACTT 558
L Q R Q L W T V L Q P S D N R L S M K L 186
TTCGGAAGCAAGAAAGGGTTACAAAAGGAAAAATATCGGCTGAGGAAGGCGGGGGTTCTT 618
F G S K K G L Q K E K Y R L R K A G V L 206
ATCATTTCATCCATGTAGTCATTTTCAGATTTTACTGGGATCTACTGATGCTGTGCCTGATC 678
I I H P C S H F R F Y W D L L M L C L I 226
ATGGCAAACGTCATCCTCCTACCCGTCGTCATTACTTTCTTCCACAACAAGGACATGAGT 738
M A N V I L L P V V I T F F H N K D M S 246
ACGGGTTGGCTCATCTTTAATTGCTTCTCAGATACCTTCTTCATTCTCGATCTCATCTGC 798
T G W L I F N C F S D T F F I L D L I C 266
AACTTTTCGGACCGGCATCATGAATCCGAAGTCGGCCGAACAGGTGATCCTCAACCCCGT 858
N F R T G I M N P K S A E Q V I L N P R 286
CAAATCGCCTATCATTATCTCCGTTTCATGGTTCATCATCGATCTCGTGTCTTCCATCCC 918
Q I A Y H Y L R S W F I I D L V S S I P 306

09640582-081700

S1

S2

S3

ATGGACTACATCTTCCTCCTCGCTGGCGGCCAGAACCGTCACTTCCTCGAGGTGTCCCGA 978
M D Y I F L L A G G Q N R H F L E V S R 326

S4

GCCCTCAAGATACTGCGCTTTGCCAAGCTCCTCAGTCTTCTTCGACTCCTGCGTCTGTCC 1038
A L K I L R F A K L L S L L R L L R L S 346

AGGCTCATGCGGTTTCGTCACTCAATGGGAACAGGCCTTCAACGTAGCCAATGCCGTCATC 1098
R L M R F V S Q W E Q A F N V A N A V I 366

S5

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R I C N L V C M M L L I G H W N G C L Q 386

TATCTCGTGCCCATGCTGCAAGAATACCCCGACCAATCATGGGTGCGCCATTAATGGCCTT 1218
Y L V P M L Q E Y P D Q S W V A I N G L 406

Pore

GAGCACGCTCATTGGTGGGAGCAGTATACATGGGCACTCTTCAAAGCCCTTTCGCACATG 1278
E H A H W W E Q Y T W A L F K A L S H M 426

CTCTGTATCGGGTACGGCAAGTTCCCCCCTCAAAGCATCACCGATGTCTGGCTAACGATT 1338
L C I G Y G K F P P Q S I T D V W L T I 446

S6

GTCAGTATGGTGTCCGGTGGGACCTGCTTCGCCCTGTTTCATCGGACACGCTACCAATCTC 1398
V S M V S G A T C F A L F I G H A T N L 466

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I Q S M D S S S R Q Y R E K L K Q V E E 486

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Y M Q Y R K L P S H L R N K I L D Y Y E 506

TACCGATACCGAGGAAAGATGTTTGATGAGAGGCATATCTTTTCGAGAAGTGTCGGAGAGT 1578
Y R Y R G K M F D E R H I F R E V S E S 526

ATACGACAGGATGTCGCAAACTACAATTGTCGCGACCTGGTTCGCATCCGTCCCTTTCTTC 1638
I R Q D V A N Y N C R D L V A S V P F F 546

GTCGGTGCCGACTCAAACCTTCGTACCCCGTGTGGTGACGCTGCTCGAATTCGAGGTCTTC 1698
V G A D S N F V T R V V T L L E F E V F 566

CAACCCGCTGACTATGTTATACAGGAAGGTACTTTTCGGTGATCGCATGTTCTTCATCCAG 1758
Q P A D Y V I Q E G T F G D R M F F I Q 586

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Q G I V D I I M S D G V I A T S L S D G 606

cNMP binding site

TCATATTTTGGCGAAATCTGCCTGCTTACCCGTGAGCGCCGCGTGGCATCGGTGAAGTGC 1878
S Y F G E I C L L T R E R R V A S V K C 626

GAGACCTACTGCACGCTCTTCTCGCTCTCCGTCCAGCATTTCAACCAAGTGCTCGACGAG 1938
E T Y C T L F S L S V Q H F N Q V L D E 646

09640582 081700

TTTCCCGCCATGAGGAAAACGATGGAAGAGATAGCCGTTCCGTCGTCTGACCCGAATCGGG	1998
F P A M R K T M E E I A V R R L T R I G	666
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K E S S K L K S R L E S P T I R D T A P	686
CTCTTTCCGATCCACCTGATACACCGTCTTTCGTCACCGACATCGAAAAGAACCGGTTT	2118
L F P I P P D T P S F V T D I E K N R F	706
TTTGGCGACGACACGGACGATGTACACATCAGGACCCGAGTCGACGTCGAGCGTGGTTTC	2178
F G D D T D D V H I R T R V D V E R G S	726
CATGAAAACGTCATCGCCATCATGGATGGGAGTTTATCCGACCTCAGGATGGAAAACGAA	2238
H E N V I A I M D G S L S D L R M E N E	746
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I Q A R K S S G K R R K F Q Q Q T T E	766
CTATGACGACTTGAAAACAAACAATGATGGACGCTTACAATTTCCAGTGATTCAATACTTA	2358
L -	767
CGCAATGCAGACATTAGCTTTTGTACCTGATTGTTTAGAATGTATTGAATTTGTAGATCA	2413
GTCCGGCAAATAAGAAAACATAAATTTGGAATTTCTTTCATTGAGGAAGTACTGAAAACAA	2478
TGTGATAGCAGCCGGTAGAAATTTCTTGTCCATTATCGAGGCTATATTTTTCGCGCTTTC	2538
TTACGAAGTAAATGAAAGGATCAATTAATTTATGTTCTTTGTCTCGTGCCTTTGTATC	2598
TGATGCCGAAAAGGAATGAAACGTGATTAGAACAGTAATCGATTGAACTACAGAAGTCTT	2658
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GAAATATTTTGTAAATTTATCTAGAATGGTACTATTGATGCTCGAAAGGTGTTGAAGTT	2778
GTCCAATATTGTGTCAAATCACCAACTATTTGACATTTGTCTTTTTC	2825

007180 28504960

Fig. 1B, 1C

B S4 motif C pore

4/18

SPIH	326- R A L K I L L R F A K L L S L L R L L R L S R L M R	350
Shaker	344- M S L A I L R V I R V F R I F K L S R H S K	368
DmEAG	341- S L F S A L K V V R L R L G R V V R K L D R Y L	365
HERG	519- E L I G L L K T A R L L R L V R V A R K L D R Y S	543
KAT I	168- S M L R L W R L R R V S S L F A R L E K D I R F N	192
brCNGC α	263- W N Y P E I R L N R L L R I S R M F E F F Q R T E	287
	416- T T A L P K A L S H M L C I G Y G K F P P Q S	438
	418- P D A F W A V V T M T T V G Y G D M T P V G	440
	441- V T A L V F T M T C M T S V G E G N V A A E T	463
	612- V T A L V F T F S S E T S V G E G N V S P N T	634
	248- V T A L V W S T T T L T T G Y G D F H A E N	270
	348- V T S L Y W S T L T L T I G - E T P P P V	368

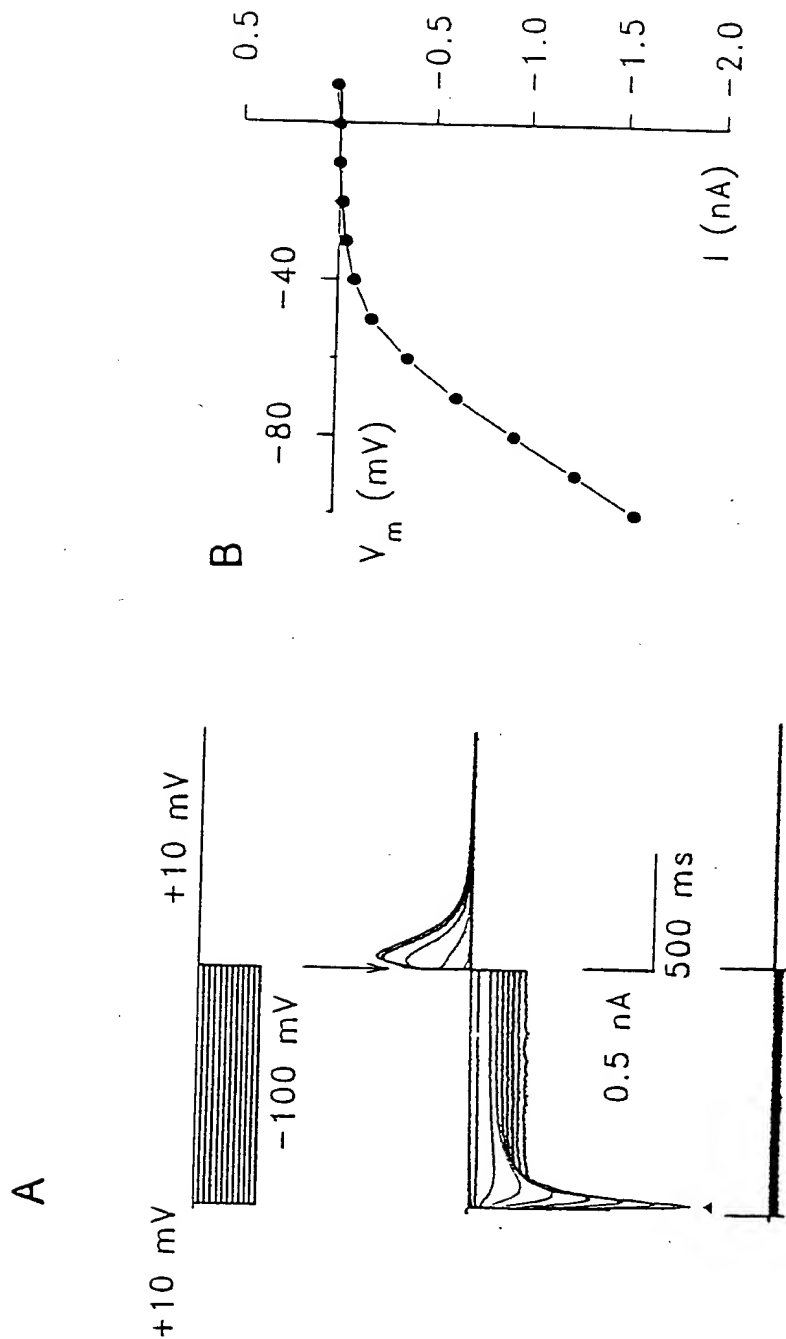
cNMP binding domain

Fig. 1D

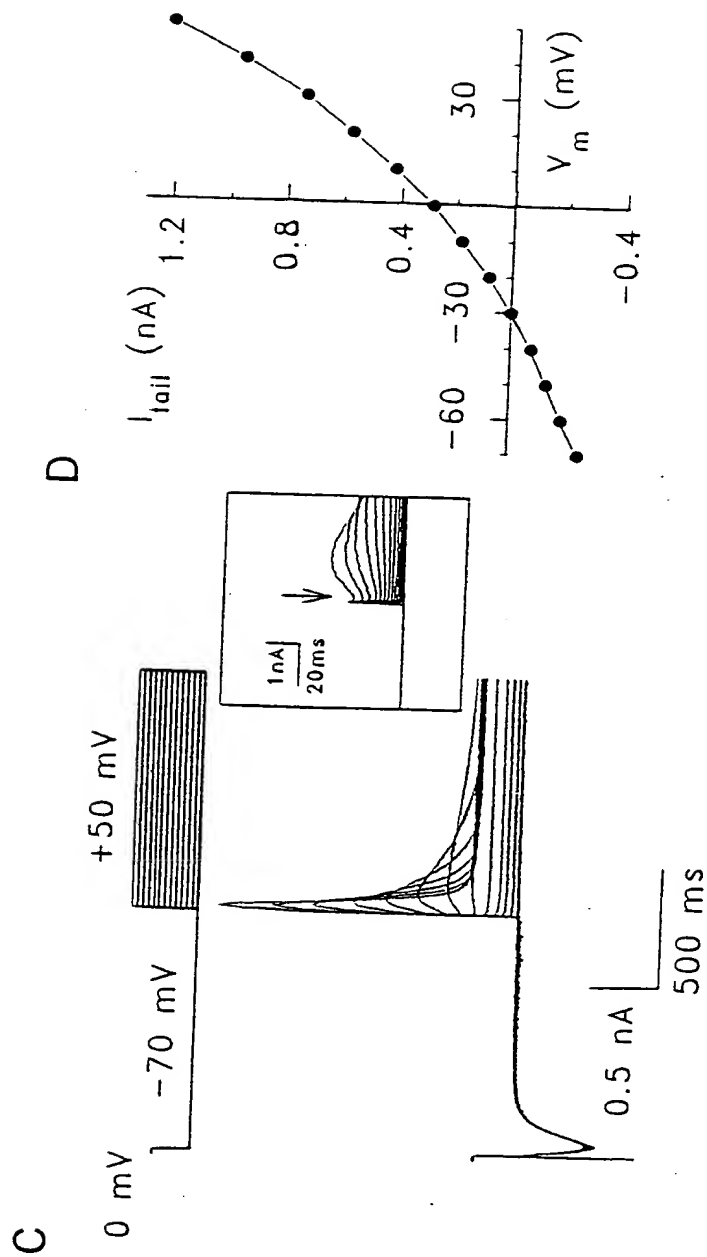
	553- FV	485- LV	462- LV	579- CL	750- CL	143- ER	110- QT	10- TE	αA	β1	β2	β3	β4	β5	β6
SPIH	UTL	VLK	VLK	AMH	AMK	FDM	VDC	EFH							
brCNGC α	VE	VE	VE	RA	RA	SD	Q	W							
boCNGC α	VE	VE	VE	RA	RA	SD	Q	W							
DmEAG	RA	RA	RA	RA	RA	SD	Q	W							
HERG	RA	RA	RA	RA	RA	SD	Q	W							
PKA I	RA	RA	RA	RA	RA	SD	Q	W							
PKG I	RA	RA	RA	RA	RA	SD	Q	W							
CAP	RA	RA	RA	RA	RA	SD	Q	W							

SPIH	C	L L	- - - -	T R E R R V A S	V K C E	T Y C T L F	S L S V Q H F N	Q V L D E S	P A M R K T M E E I A	V R R L T P I G A E S S	* -670
b r C N G C α	S	I L N I K G S K A	G R R R T A N I	K S I G Y S D L F C	L S K D	D L M B A L T E	P D A K G M E E E K G K	Q I L M R D G L L D I			-609
b o C N G C α	S	I L N I K G S K M G	R R R T A N I	S L G Y S D L F C	L S K D	D L M B A T E	P D A K R V L E E R G R E I L M K E G	L L D E			-586
D m E A G	F W K D S	- - - -	A V G Q S A A N	V F A L T Y C D L H A	A T K R D X L L E	V L D F Y S A F A N S	F A R N L V L T Y N L R H R	L I F			-697
H E R G	L N L Y A R	- - - -	P G K S N G	D V R A L T Y C D L H K	H R D D L L E	V L D M Y P E F S D H F W S S L E	I T F N L R D T N M I				-868
P K A I	A L L Y	- - - -	G T P R A A	T V K A K T N V K L N G	H D R S	R R L L M G S T	L R R K M Y E E F F S K V S I L	E S L D -			-258
P K G I	A L L Y N	- - - -	C T R T A T	V K T L V N V K L W S I	D R O C F C T	I M M R T G L I K H T E Y M E F F K S V P T F Q S L P E					-226
C A P	G L F E E	- - - -	G O E R S A W	R A K T A C E V A E I S Y K I K F R Q	H I Q V N P D I L M R H	S A Q M A R R L Q V T S E K V G					-132

Fig. 2A, 2B



Fig, 2C, 2D



Fig, 2E, 2F

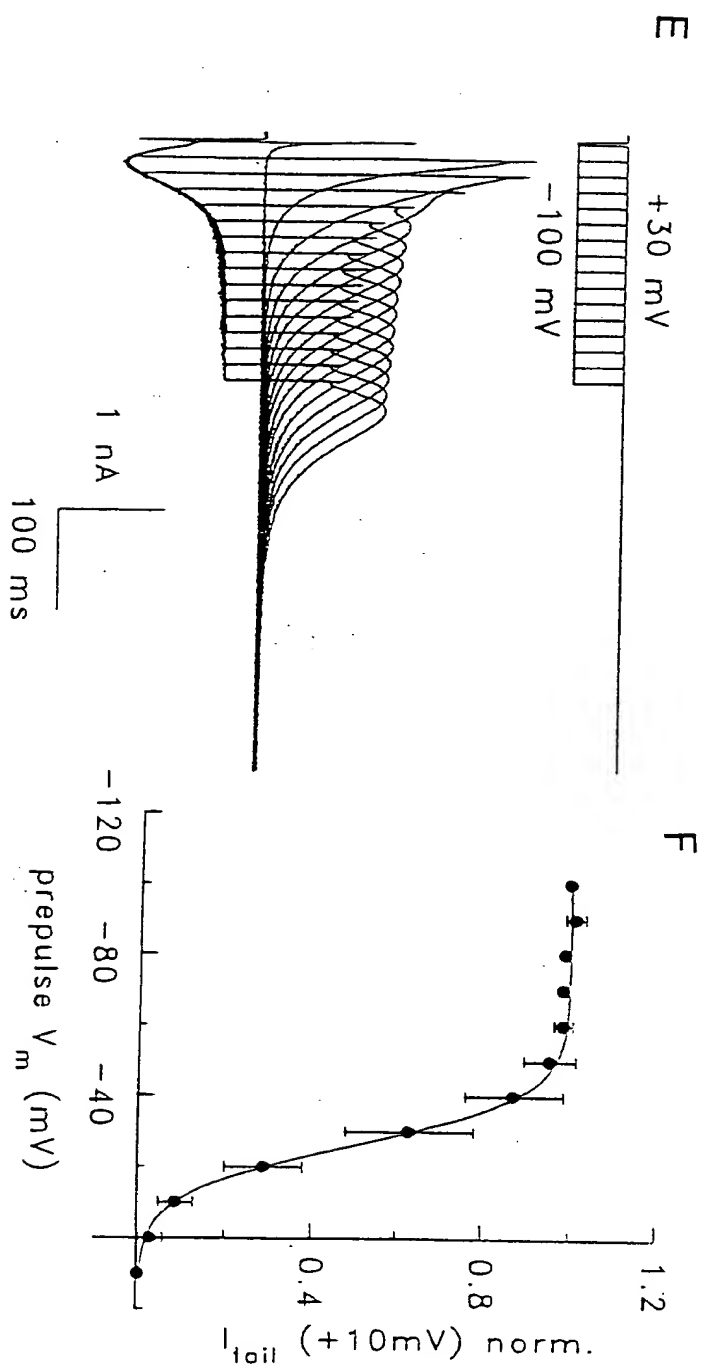


Fig. 3A, 3B

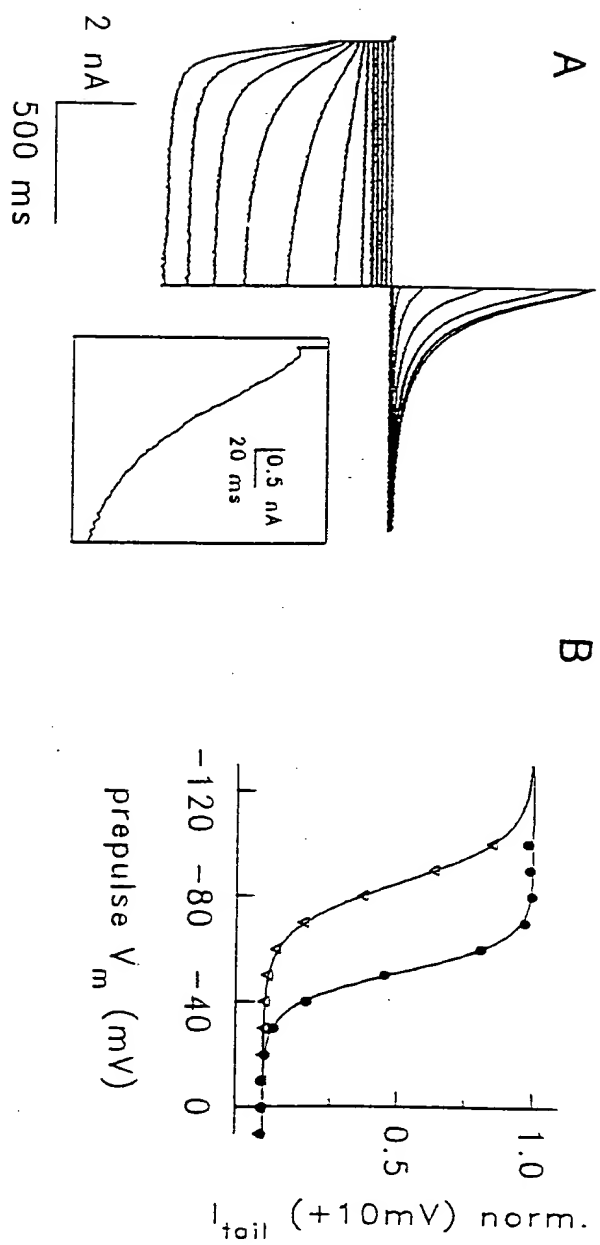


Fig. 3C, 3D

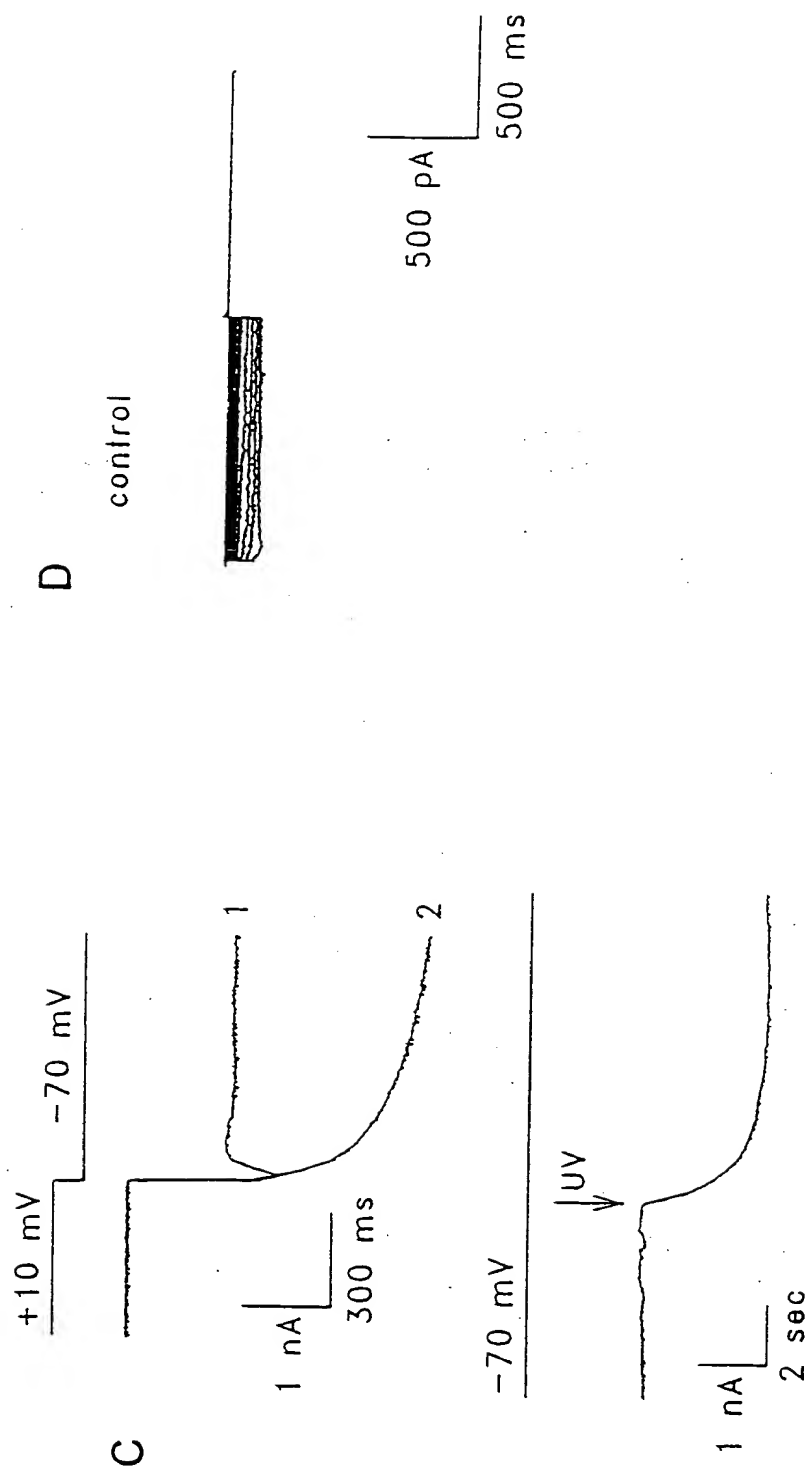
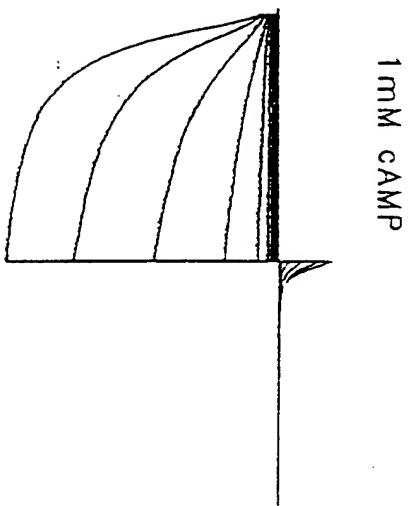


Fig. 3E, 3F

E



F

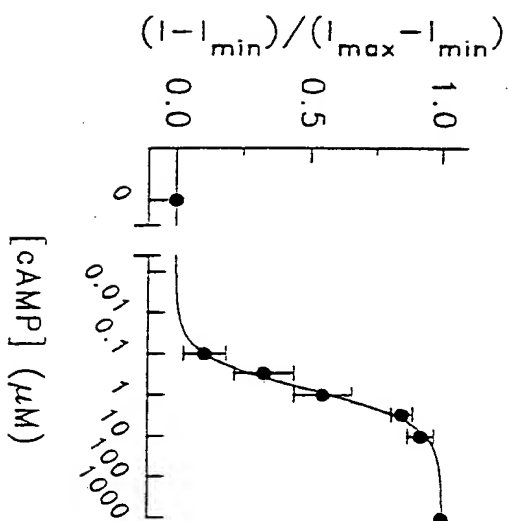


Fig. 4

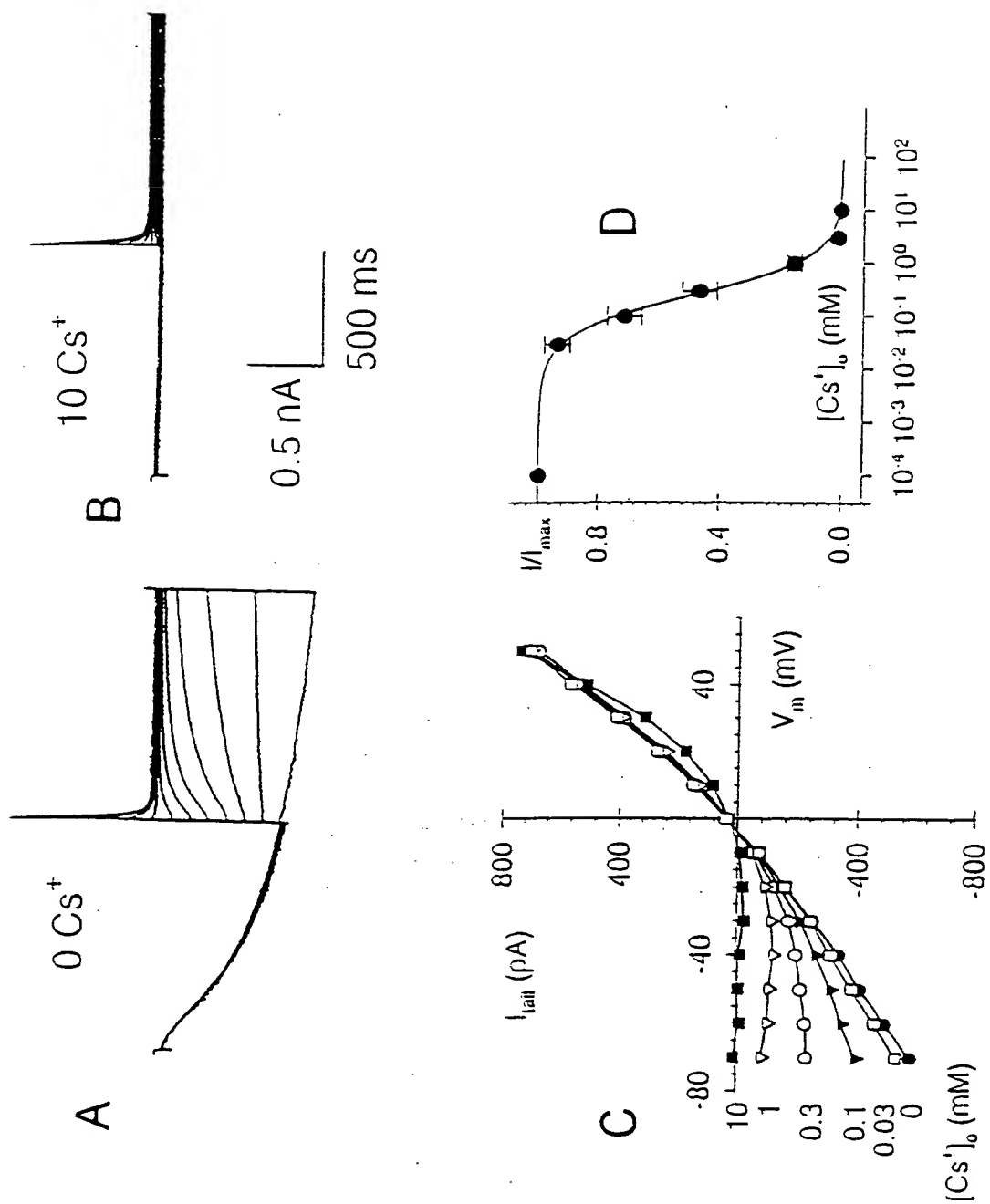


Fig. 4E

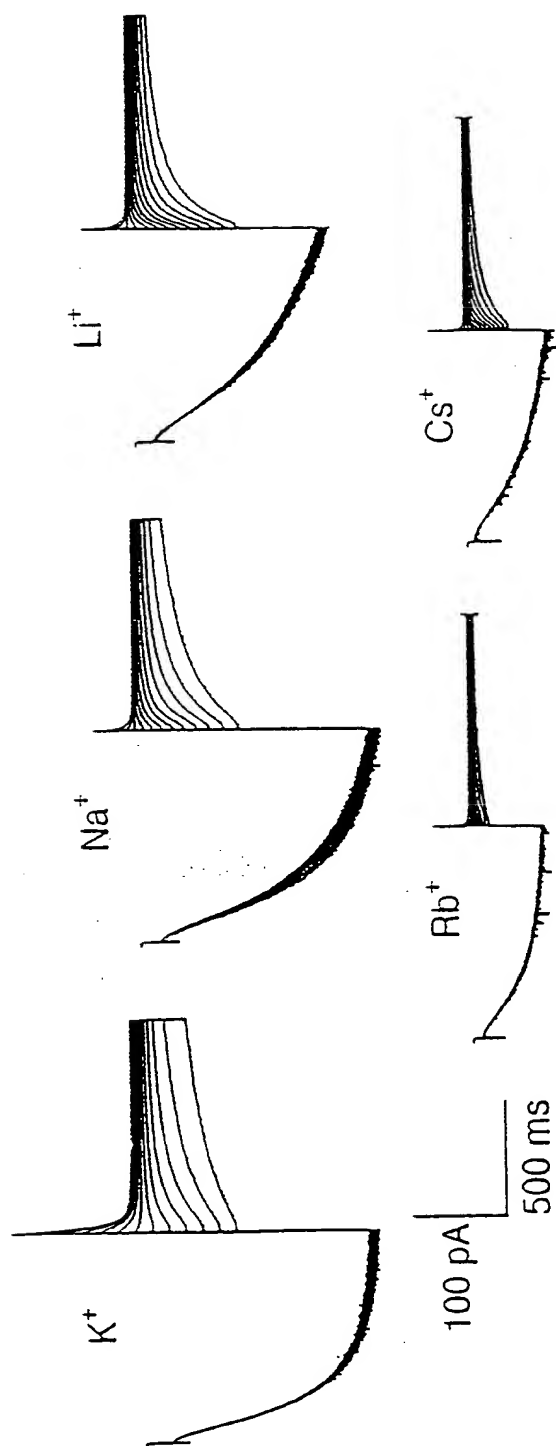


Fig. 4F

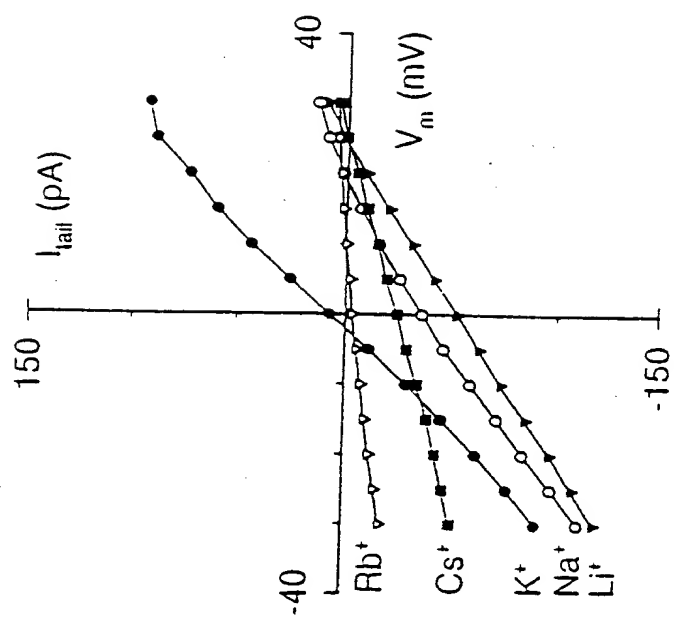


Fig. 4G

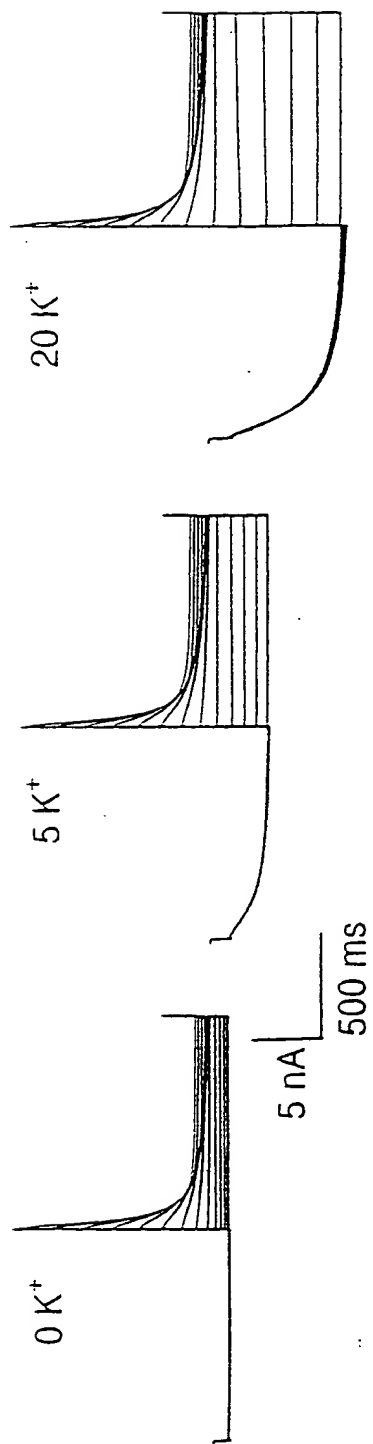


Fig. 4H

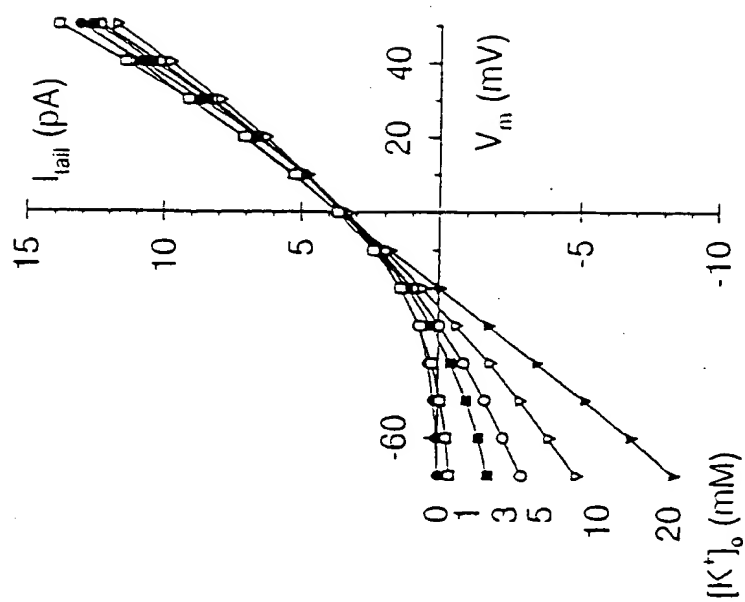


Fig. 5

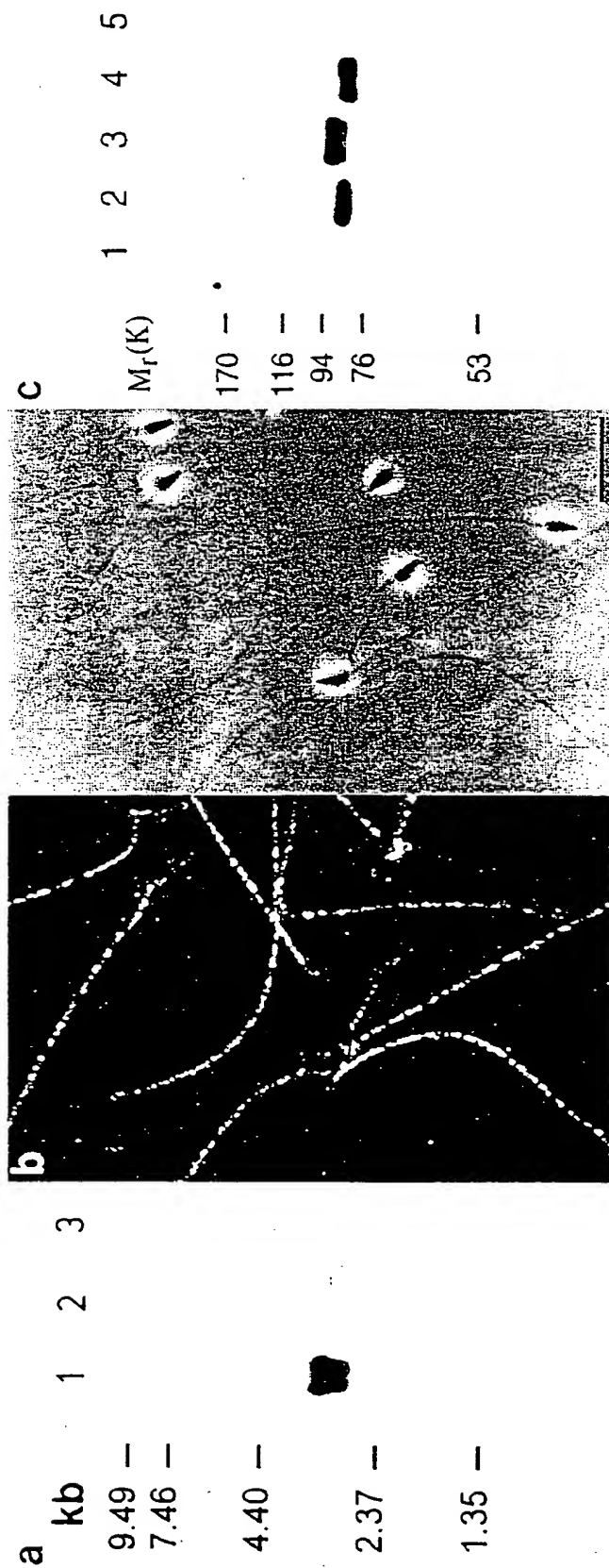


Fig. 6

